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1 SEQUENCE LISTING

<110> Sema ApS

Albrechtsen, Morten

Christensen, Claus

Lukanidin, Eugene

Olesen, Ole

<120> Use of compounds capable of inhibiting the proteolytic processing of semaphorins for prevention, treatment, diagnosis and prognosis of an invasive disease

<130> P682 PC00

<160> 30

<170> PatentIn version 3.1

<210> 1

<211> 775

<212> PRT

<213> mouse Sema3E Genbank#Z93948

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Leu Arg Leu Ser His Lys Glu Leu Leu Glu Leu Asn Arg Thr Ser Ile 35 40 45

Phe Gln Ser Pro Leu Gly Phe Leu Asp Leu His Thr Met Leu Leu Asp 50 55 60

Glu Tyr Gln Glu Arg Leu Phe Val Gly Gly Arg Asp Leu Val Tyr Ser 65 70 75 80

Leu Asn Leu Glu Arg Val Ser Asp Gly Tyr Arg Glu Ile Tyr Trp Pro 85 90 95

Ser Thr Ala Val Lys Val Glu Glu Cys Ile Met Lys Gly Lys Asp Ala 100 105 110

Asn Glu Cys Ala Asn Tyr Ile Arg Val Leu His His Tyr Asn Arg Thr 115 120 125

His Leu Leu Thr Cys Ala Thr Gly Ala Phe Asp Pro His Cys Ala Phe 130 135 140

Ile Arg Val Gly His His Ser Glu Glu Pro Leu Phe His Leu Glu Ser 145 150 155 160

His Arg Ser Glu Arg Gly Arg Gly Arg Cys Pro Phe Asp Pro Asn Ser 165 170 175

Ser Phe Val Ser Thr Leu Val Gly Asn Glu Leu Phe Ala Gly Leu Tyr 180 185 190

Ser Asp Tyr Trp Gly Arg Asp Ser Ala Ile Phe Arg Ser Met Gly Lys 195 200 205

Leu Gly His Ile Arg Thr Glu His Asp Asp Glu Arg Leu Leu Lys Glu 210 215 220

Pro Lys Phe Val Gly Ser Tyr Met Ile Pro Asp Asn Glu Asp Arg Asp 225 230 235 240

Asp Asn Lys Met Tyr Phe Phe Phe Thr Glu Lys Ala Leu Glu Ala Glu 250 245 Asn Asn Ala His Thr Ile Tyr Thr Arg Val Gly Arg Leu Cys Val Asn Asp Met Gly Gly Gln Arg Ile Leu Val Asn Lys Trp Ser Thr Phe Leu Lys Ala Arg Leu Val Cys Ser Val Pro Gly Met Asn Gly Ile Asp Thr 290 Tyr Phe Asp Glu Leu Glu Asp Val Phe Leu Leu Pro Thr Arg Asp Pro 310 315 305 Lys Asn Pro Val Ile Phe Gly Leu Phe Asn Thr Thr Ser Asn Ile Phe 335 330 325 Arg Gly His Ala Val Cys Val Tyr His Met Ser Ser Ile Arg Glu Ala 345 350 340 Phe Asn Gly Pro Tyr Ala His Lys Glu Gly Pro Glu Tyr His Trp Ser 360 355 Leu Tyr Glu Gly Lys Val Pro Tyr Pro Arg Pro Gly Ser Cys Ala Ser 380 375 Lys Val Asn Gly Gly Lys Tyr Gly Thr Thr Lys Asp Tyr Pro Asp Asp Ala Ile Arg Phe Ala Arg Ile Asp Pro Leu Met Tyr Gln Pro Ile Lys 405 Pro Val His Lys Lys Pro Ile Leu Val Lys Thr Asp Gly Lys Tyr Asn 425 420 Leu Arg Gln Leu Ala Val Asp Arg Val Glu Ala Glu Asp Gly Gln Tyr 440 445 435 Asp Val Leu Phe Ile Gly Thr Asp Thr Gly Ile Val Leu Lys Val Ile 450 455 460

Thr Ile Tyr Asn Gln Glu Thr Glu Trp Met Glu Glu Val Ile Leu Glu Glu Leu Gln Ile Phe Lys Asp Pro Ala Pro Ile Ile Ser Met Glu Ile Ser Ser Lys Arg Gln Gln Leu Tyr Ile Gly Ser Ala Ser Ala Val Ala Gln Val Arg Phe His His Cys Asp Met Tyr Gly Ser Ala Cys Ala Asp Cys Cys Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys Ser Arg Tyr Tyr Pro Thr Gly Ala His Ala Lys Arg Arg Phe Arg Arg Gln Asp Val Arg His Gly Asn Ala Ala Gln Gln Cys Phe Gly Gln Gln Phe Val Gly Asp Ala Leu Asp Arg Thr Glu Glu Arg Leu Ala Tyr Gly Ile Glu Ser Asn Ser Thr Leu Leu Glu Cys Thr Pro Arg Ser Leu Gln Ala Lys Val Ile Trp Phe Tyr Gln Lys Gly Arg Asp Val Arg Lys Glu Glu Val Lys Thr Asp Asp Arg Val Val Lys Met Asp Leu Gly Leu Leu Phe Leu Arg Val Arg Lys Ser Asp Ala Gly Thr Tyr Phe Cys Gln Thr

Val Glu His Asn Phe Val His Thr Val Arg Lys Ile Thr Leu Glu Val

Val Glu Glu His Lys Val Glu Gly Met Phe His Lys Asp His Glu Glu 675 680 685

Glu Arg His His Lys Met Pro Cys Pro Pro Leu Ser Gly Met Ser Gln 690 695 700

Gly Thr Lys Pro Trp Tyr Lys Glu Phe Leu Gln Leu Ile Gly Tyr Ser 705 710 715 720

Asn Phe Gln Arg Val Glu Glu Tyr Cys Glu Lys Val Trp Cys Thr Asp 725 730 735

Lys Lys Arg Lys Lys Leu Lys Met Ser Pro Ser Lys Trp Lys Tyr Ala 740 745 750

Asn Pro Gln Glu Lys Arg Leu Arg Ser Lys Ala Glu His Phe Arg Leu 755 760 765

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Leu Arg Leu Ser His Lys Glu Leu Leu Asn Leu Asn Arg Thr Ser Ile 35 40 45

Phe His Ser Pro Phe Gly Phe Leu Asp Leu His Thr Met Leu Leu Asp 50 55 60

Glu Tyr Gln Glu Arg Leu Phe Val Gly Gly Arg Asp Leu Val Tyr Ser 65 70 75 80

Leu Ser Leu Glu Arg Ile Ser Asp Gly Tyr Lys Glu Ile His Trp Pro 85 90 95

Ser Thr Ala Leu Lys Met Glu Glu Cys Ile Met Lys Gly Lys Asp Ala 100 105 110

Gly Glu Cys Ala Asn Tyr Val Arg Val Leu His His Tyr Asn Arg Thr 115 120 125

His Leu Leu Thr Cys Gly Thr Gly Ala Phe Asp Pro Val Cys Ala Phe 130 135 140

Ile Arg Val Gly Tyr His Leu Glu Asp Pro Leu Phe His Leu Glu Ser 145 150 155 160

Pro Arg Ser Glu Arg Gly Arg Gly Arg Cys Pro Phe Asp Pro Ser Ser 165 170 175

Ser Phe Ile Ser Thr Leu Ile Gly Ser Glu Leu Phe Ala Gly Leu Tyr 180 185 190

Ser Asp Tyr Trp Ser Arg Asp Ala Ala Ile Phe Arg Ser Met Gly Arg 195 200 205

Leu Ala His Ile Arg Thr Glu His Asp Asp Glu Arg Leu Leu Lys Glu 210 215 220

Pro Lys Phe Val Gly Ser Tyr Met Ile Pro Asp Asn Glu Asp Arg Asp 225 230 235 240

Asp Asn Lys Val Tyr Phe Phe Phe Thr Glu Lys Ala Leu Glu Ala Glu 245 250 255

Asn Asn Ala His Ala Ile Tyr Thr Arg Val Gly Arg Leu Cys Val Asn Asp Val Gly Gly Gln Arg Ile Leu Val Asn Lys Trp Ser Thr Phe Leu Lys Ala Arg Leu Val Cys Ser Val Pro Gly Met Asn Gly Ile Asp Thr Tyr Phe Asp Glu Leu Glu Asp Val Phe Leu Leu Pro Thr Arg Asp His Lys Asn Pro Val Ile Phe Gly Leu Phe Asn Thr Thr Ser Asn Ile Phe Arg Gly His Ala Ile Cys Val Tyr His Met Ser Ser Ile Arg Ala Ala Phe Asn Gly Pro Tyr Ala His Lys Glu Gly Pro Glu Tyr His Trp Ser Val Tyr Glu Gly Lys Val Pro Tyr Pro Arg Pro Gly Ser Cys Ala Ser Lys Val Asn Gly Gly Arg Tyr Gly Thr Thr Lys Asp Tyr Pro Asp Asp Ala Ile Arg Phe Ala Arg Ser His Pro Leu Met Tyr Gln Ala Ile Lys Pro Ala His Lys Lys Pro Ile Leu Val Lys Thr Asp Gly Lys Tyr Asn Leu Lys Gln Ile Ala Val Asp Arg Val Glu Ala Glu Asp Gly Gln Tyr Asp Val Leu Phe Ile Gly Thr Asp Asn Gly Ile Val Leu Lys Val Ile Thr Ile Tyr Asn Gln Glu Met Glu Ser Met Glu Glu Val Ile Leu Glu 465 470 475 480

Glu Leu Gln Ile Phe Lys Asp Pro Val Pro Ile Ile Ser Met Glu Ile 485 490 495

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Gln Val Arg Phe His His Cys Asp Met Tyr Gly Ser Ala Cys Ala Asp 515 520 525

Cys Cys Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys 530 540

Ser Arg Tyr Tyr Pro Thr Gly Thr His Ala Lys Arg Arg Phe Arg Arg 545 550 550 560

Gln Asp Val Arg His Gly Asn Ala Ala Gln Gln Cys Phe Gly Gln Gln 565 570 575

Phe Val Gly Asp Ala Leu Asp Lys Thr Glu Glu His Leu Ala Tyr Gly 580 585 590

Ile Glu Asn Asn Ser Thr Leu Leu Glu Cys Thr Pro Arg Ser Leu Gln 595 600 605

Ala Lys Val Ile Trp Phe Val Gln Lys Gly Arg Glu Thr Arg Lys Glu 610 615 620

Glu Val Lys Thr Asp Asp Arg Val Val Lys Met Asp Leu Gly Leu Leu 625 630 635 640

Phe Leu Arg Leu His Lys Ser Asp Ala Gly Thr Tyr Phe Cys Gln Thr 645 650 655

Val Glu His Ser Phe Val His Thr Val Arg Lys Ile Thr Leu Glu Val 660 665 670

Val Glu Glu Lys Val Glu Asp Met Phe Asn Lys Asp Asp Glu Glu 675 680 685

Asp Arg His His Arg Met Pro Cys Pro Ala Gln Ser Ser Ile Ser Gln 690 695 700

Gly Ala Lys Pro Trp Tyr Lys Glu Phe Leu Gln Leu Ile Gly Tyr Ser 705 710 715 720

Asn Phe Gln Arg Val Glu Glu Tyr Cys Glu Lys Val Trp Cys Thr Asp 725 730 735

Arg Lys Arg Lys Lys Leu Lys Met Ser Pro Ser Lys Trp Lys Tyr Ala 740 745 750

Asn Pro Gln Glu Lys Lys Leu Arg Ser Lys Pro Glu His Tyr Arg Leu 755 760 765

Pro Arg His Thr Leu Asp Ser 770 775

<210> 3

<211> 560

<212> PRT

<213> 61 kDa proteolytic fragment of mouse Sema3E

<400> 3

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Leu Glu Leu Trp Thr Pro Gly His Ser Ala Asn Pro Ser Tyr Pro Arg 20 25 30

Leu Arg Leu Ser His Lys Glu Leu Leu Glu Leu Asn Arg Thr Ser Ile 35 40 45

Phe Gln Ser Pro Leu Gly Phe Leu Asp Leu His Thr Met Leu Leu Asp 50 55 60

Glu Tyr Gln Glu Arg Leu Phe Val Gly Gly Arg Asp Leu Val Tyr Ser 65 70 75 80

Leu Asn Leu Glu Arg Val Ser Asp Gly Tyr Arg Glu Ile Tyr Trp Pro 85 90 95

Ser Thr Ala Val Lys Val Glu Glu Cys Ile Met Lys Gly Lys Asp Ala 100 105 110

Asn Glu Cys Ala Asn Tyr Ile Arg Val Leu His His Tyr Asn Arg Thr 115 120 125

His Leu Leu Thr Cys Ala Thr Gly Ala Phe Asp Pro His Cys Ala Phe 130 135 140

Ile Arg Val Gly His His Ser Glu Glu Pro Leu Phe His Leu Glu Ser 145 150 155 160

His Arg Ser Glu Arg Gly Arg Gly Arg Cys Pro Phe Asp Pro Asn Ser 165 170 175

Ser Phe Val Ser Thr Leu Val Gly Asn Glu Leu Phe Ala Gly Leu Tyr 180 185 190

Ser Asp Tyr Trp Gly Arg Asp Ser Ala Ile Phe Arg Ser Met Gly Lys 195 200 205

Leu Gly His Ile Arg Thr Glu His Asp Asp Glu Arg Leu Leu Lys Glu 210 215 220

Pro Lys Phe Val Gly Ser Tyr Met Ile Pro Asp Asn Glu Asp Arg Asp 225 230 235

Asp Asn Lys Met Tyr Phe Phe Phe Thr Glu Lys Ala Leu Glu Ala Glu 245 250 255

Asn Asn Ala His Thr Ile Tyr Thr Arg Val Gly Arg Leu Cys Val Asn 260 265 270

Asp Met Gly Gly Gln Arg Ile Leu Val Asn Lys Trp Ser Thr Phe Leu Lys Ala Arg Leu Val Cys Ser Val Pro Gly Met Asn Gly Ile Asp Thr Tyr Phe Asp Glu Leu Glu Asp Val Phe Leu Leu Pro Thr Arg Asp Pro Lys Asn Pro Val Ile Phe Gly Leu Phe Asn Thr Thr Ser Asn Ile Phe Arg Gly His Ala Val Cys Val Tyr His Met Ser Ser Ile Arg Glu Ala Phe Asn Gly Pro Tyr Ala His Lys Glu Gly Pro Glu Tyr His Trp Ser Leu Tyr Glu Gly Lys Val Pro Tyr Pro Arg Pro Gly Ser Cys Ala Ser Lys Val Asn Gly Gly Lys Tyr Gly Thr Thr Lys Asp Tyr Pro Asp Asp Ala Ile Arg Phe Ala Arg Ile Asp Pro Leu Met Tyr Gln Pro Ile Lys Pro Val His Lys Lys Pro Ile Leu Val Lys Thr Asp Gly Lys Tyr Asn Leu Arg Gln Leu Ala Val Asp Arg Val Glu Ala Glu Asp Gly Gln Tyr Asp Val Leu Phe Ile Gly Thr Asp Thr Gly Ile Val Leu Lys Val Ile Thr Ile Tyr Asn Gln Glu Thr Glu Trp Met Glu Glu Val Ile Leu Glu Glu Leu Gln Ile Phe Lys Asp Pro Ala Pro Ile Ile Ser Met Glu Ile

12 485 490

495

Ser Ser Lys Arg Gln Gln Leu Tyr Ile Gly Ser Ala Ser Ala Val Ala 500 505 510

Gln Val Arg Phe His His Cys Asp Met Tyr Gly Ser Ala Cys Ala Asp 515 520 525

Cys Cys Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys 530 540

Ser Arg Tyr Tyr Pro Thr Gly Ala His Ala Lys Arg Arg Phe Arg Arg 545 550 555 560

<210> 4

<211> 560

<212> PRT

<213> 61 kDa proteolytic fragment of human SEMA3E

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Leu Glu Leu Trp Thr Gly Gly His Thr Ala Asp Thr Thr His Pro Arg
20 25 30

Leu Arg Leu Ser His Lys Glu Leu Leu Asn Leu Asn Arg Thr Ser Ile 35 40 45

Phe His Ser Pro Phe Gly Phe Leu Asp Leu His Thr Met Leu Leu Asp 50 55 60

Glu Tyr Gln Glu Arg Leu Phe Val Gly Gly Arg Asp Leu Val Tyr Ser 65 70 75 80

Leu Ser Leu Glu Arg Ile Ser Asp Gly Tyr Lys Glu Ile His Trp Pro 85 90 95

13

Ser Thr Ala Leu Lys Met Glu Glu Cys Ile Met Lys Gly Lys Asp Ala 100 105 110

Gly Glu Cys Ala Asn Tyr Val Arg Val Leu His His Tyr Asn Arg Thr 115 120 125

His Leu Leu Thr Cys Gly Thr Gly Ala Phe Asp Pro Val Cys Ala Phe 130 135 140

Ile Arg Val Gly Tyr His Leu Glu Asp Pro Leu Phe His Leu Glu Ser 145 150 155 160

Pro Arg Ser Glu Arg Gly Arg Gly Arg Cys Pro Phe Asp Pro Ser Ser 165 170 175

Ser Phe Ile Ser Thr Leu Ile Gly Ser Glu Leu Phe Ala Gly Leu Tyr 180 185 190

Ser Asp Tyr Trp Ser Arg Asp Ala Ala Ile Phe Arg Ser Met Gly Arg 195 200 205

Leu Ala His Ile Arg Thr Glu His Asp Asp Glu Arg Leu Leu Lys Glu 210 215 220

Pro Lys Phe Val Gly Ser Tyr Met Ile Pro Asp Asn Glu Asp Arg Asp 225 230 235 240

Asp Asn Lys Val Tyr Phe Phe Phe Thr Glu Lys Ala Leu Glu Ala Glu 245 250 255

Asn Asn Ala His Ala Ile Tyr Thr Arg Val Gly Arg Leu Cys Val Asn 260 265 270

Asp Val Gly Gly Gln Arg Ile Leu Val Asn Lys Trp Ser Thr Phe Leu 275 280 285

Lys Ala Arg Leu Val Cys Ser Val Pro Gly Met Asn Gly Ile Asp Thr 290 295 300

14

Tyr Phe Asp Glu Leu Glu Asp Val Phe Leu Leu Pro Thr Arg Asp His 305 310 315 320

Lys Asn Pro Val Ile Phe Gly Leu Phe Asn Thr Thr Ser Asn Ile Phe 325 330 335

Arg Gly His Ala Ile Cys Val Tyr His Met Ser Ser Ile Arg Ala Ala 340 345 350

Phe Asn Gly Pro Tyr Ala His Lys Glu Gly Pro Glu Tyr His Trp Ser 355 360 365

Val Tyr Glu Gly Lys Val Pro Tyr Pro Arg Pro Gly Ser Cys Ala Ser 370 375 380

Lys Val Asn Gly Gly Arg Tyr Gly Thr Thr Lys Asp Tyr Pro Asp Asp 385 390 395 400

Ala Ile Arg Phe Ala Arg Ser His Pro Leu Met Tyr Gln Ala Ile Lys 405 410 415

Pro Ala His Lys Lys Pro Ile Leu Val Lys Thr Asp Gly Lys Tyr Asn 420 425 430

Leu Lys Gln Ile Ala Val Asp Arg Val Glu Ala Glu Asp Gly Gln Tyr 435 440 445

Asp Val Leu Phe Ile Gly Thr Asp Asn Gly Ile Val Leu Lys Val Ile 450 455 460

Thr Ile Tyr Asn Gln Glu Met Glu Ser Met Glu Glu Val Ile Leu Glu 465 470 475 480

Glu Leu Gln Ile Phe Lys Asp Pro Val Pro Ile Ile Ser Met Glu Ile 485 490 495

Ser Ser Lys Arg Gln Gln Leu Tyr Ile Gly Ser Ala Ser Ala Val Ala 500 505 510

15

Gln Val Arg Phe His His Cys Asp Met Tyr Gly Ser Ala Cys Ala Asp 515 520 525

Cys Cys Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys 530 535 540

Ser Arg Tyr Tyr Pro Thr Gly Thr His Ala Lys Arg Arg Phe Arg Arg 545 550 555 560

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<211> 50

<212> PRT

<213> mouse Sema3E : a fragment comprising a pro-protein
convertase cleavage site

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Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys Ser Arg 1 5 10 15

Tyr Tyr Pro Thr Gly Thr His Ala Lys Arg Arg Phe Arg Arg Gln Asp 20 25 30

Val Arg His Gly Asn Ala Ala Gln Gln Cys Phe Gly Gln Gln Phe Val 35 40 45

Gly Asp 50

<210> 6

<211> 50

<212> PRT

<213> human SEMA3E: a fragment comprising a pro-protein convertase cleavage site

<400> 6

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16

Leu Ala Arg Asp Pro Tyr Cys Ala Trp Asp Gly Ile Ser Cys Ser Arg 1 5 10 15

Tyr Tyr Pro Thr Gly Thr His Ala Lys Arg Arg Phe Arg Arg Gln Asp 20 25 30

Val Arg His Gly Asn Ala Ala Gln Gln Cys Phe Gly Gln Gln Phe Val 35 40 45

Gly Asp 50

<210> 7

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<212> DNA

<213> cDNA mouse Sema3E #Z93947

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cagcetecte etteacteeg egtetggget gaeggegaea geaceageee ggaeetgget 360

ctcaagacgc gctccttgga cggtctcttg ctccgcgctt ctaaccaccg ggcccaaaga 420

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540

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18

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23

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27

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Cys Leu Lys Ala Asp Pro Arg Phe Glu Cys Gly Trp Cys Val Ala Glu 260 265 270

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Leu Lys Leu Ser Pro Glu Thr Gly Pro Arg Gln Gly Gly Thr Arg Leu 305 310 315 320

Thr Ile Thr Gly Glu Asn Leu Gly Leu Arg Phe Glu Asp Val Arg Leu 325 330 335

Gly Val Arg Val Gly Lys Val Leu Cys Ser Pro Val Glu Ser Glu Tyr 340 345 350

Ile Ser Ala Glu Gln Ile Val Cys Glu Ile Gly Asp Ala Ser Ser Val 355 360 365

Arg Ala His Asp Ala Leu Val Glu Val Cys Val Arg Asp Cys Ser Pro 370 375 380

His Tyr Arg Ala Leu Ser Pro Lys Arg Phe Thr Phe Val Thr Pro Thr 385 390 395 400

Phe Tyr Arg Val Ser Pro Ser Arg Gly Pro Leu Ser Gly Gly Thr Trp 405 410 415

Ile Gly Ile Glu Gly Ser His Leu Asn Ala Gly Ser Asp Val Ala Val 420 425 430

Ser Val Gly Gly Arg Pro Cys Ser Phe Ser Trp Arg Asn Ser Arg Glu 435 440 445

Ile Arg Cys Leu Thr Pro Pro Gly Gln Ser Pro Gly Ser Ala Pro Ile 450 455 460

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Ser Gly Ser Gln Val Ile Cys Ile Ser Pro Gly Pro Lys Asp Val Pro 50 55 60

Val Ile Pro Leu Asp Gln Asp Trp Phe Gly Leu Glu Leu Gln Leu Arg 65 70 75 80

Ser Lys Glu Thr Gly Lys Ile Phe Val Ser Thr Glu Phe Lys Phe Tyr 85 90 95

Asn Cys Ser Ala His Gln Leu Cys Leu Ser Cys Val Asn Ser Ala Phe 100 105 110

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Gln Leu Val Pro Thr Glu Glu Ile Leu Ile Pro Val Gly Glu Val Lys

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34

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Thr Gly His Tyr Leu Gly Ala Gly Ser Ser Val Ala Val Tyr Leu Gly 420 425 430

Asn Gln Thr Cys Glu Phe Tyr Gly Arg Ser Met Ser Glu Ile Val Cys 435 440 445

Val Ser Pro Pro Ser Ser Asn Gly Leu Gly Pro Val Pro Val Ser Val 450 455 460

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Gly His Thr Pro Leu Thr Ile Thr Gly Phe Asn Leu Asp Val Ile Gln 500 510

Glu Pro Arg Ile Arg Val Lys Phe Asn Gly Lys Glu Ser Val Asn Val 515 520 525

Cys Lys Val Val Asn Thr Thr Leu Thr Cys Leu Ala Pro Ser Leu 530 535 540

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Phe Gly Phe Val Phe Asn Asn Val Gln Ser Leu Leu Ile Tyr Asn Asp 565 570 575

35

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